αv	β8	C6D4 VH	C6D4 VL
E121			R33
D148			R33
D218			R35
	H118		Y55
	D171		W56
	N173		K95
	N173		S97
	D175	S33	
	D175	R50	
	K203	D31	

Table S1 Interactions of αvβ8 with Fab C6D4, related to Fig. 2

αv	β8	C6-RGD3 VH	C6-RGD3 VL
D148			R32
F177			R37
	D171		W60
	N173		K99
	N173		S101
	N173	Y101	
	R201	T28	
	Q202	R103	
	K203	D31	
	I204	Y100	
	Mg2+ (MIDAS)		D34

Table S2 Interactions of αvβ8 with Fab C6-RGD3, related to Fig. 2

		W2				W3				W3				W4
C6-RGD3						L_1				L_1				
C6D4		L_1				L_1								L_1
αv	119	KQEREP		αv	146	DIDADG		αv	175	GSFYWQ		αv	210	TRTAQAIFDDSY
α8	124	TPEKDP		α8	153	NADPEG		α8	182	GSFYWQ		α8	217	TEVAPASYDDSY
α5	126	EPLSDP		α5	155	FSWAAG		α5	184	GSYFWQ		α5	219	TRQASSIYDDSY
α_{iib}	121	EAEKTP		α_{iib}	160	FSWDKR		α_{iib}	187	GGYYFL		α_{iib}	223	FDSSNPEYFDGY
		=								=				= =

αv subunit interactions with L-TGF-β, C6D4, and C6-RGD3
= conserved

Table S3 C6D4, C6-RGD3, L-TGF-β1 interactions with αv subunit β-propeller blades W2, W3, W4 aligned to other α-subunits, related to Fig. 2

pL-TGF-β1[a]	192	AHCSCDSKDNT----------LHVEING---FNSGRRGDLATIHG------MNRP
hL-TGF-β1[b]	192	AHCSCDSRDNT----------LQVDING---FTTGRRGDLATIHG------MNRP
hL-TGF-β3	202	IHCPCHTFQPNGD-ILENIHEVMEIKFKGVDNEDDHGRGDLGRLKK---QKDHHNP
hL-TGF-β2	204	LHCPCCTFVPSNNYIIPNKSEELEARFAGIDGTSTYTSGDQKTIKSTRKKNSGKTP
		== = = == =
C6D4 CDR L_1	30	L-NSRTRK---N
C6-RGD3 CDR L_1	30	LGRGDLGRLKKN
		= =

interactions between αvβ8/L-TGF-β, αvβ8/C6D4, αvβ8/C6-RGD3
hydrophobic interactions between αvβ8/L-TGF-β
[a]porcine L-TGF-β1
[b]human L-TGF-β1
= conserved

Table S4 C6D4 and C6-RGD3 CDRL1 loop alignment to TGF-β isoforms, related to Fig.2

Primer Name	Sequence
C6-RGD3 CDRL1 loop F	5'-GATCTGGGGCGCCTCAAGAAGAACGCCTTGGCTTGGTACCAGCAG-3'
C6-RGD3 CDRL1 loop R	5'-CTTGAGGCGCCCCAGATCTCCACGGCCGAGCAGACTCTGACTGGATTTG-3'
β8 I208R F	5' CAGAAGATCTCTGGAAACAGAGATACACC-3'
β8 I208R R	5'-GAAGTTTGGTCGACATAATGC-3'
β8 Y172N F	5'-CAATGCAGTGACAACAATTTAGACTGC-3'
β8 Y172N R	5'-GCAGTCTAAATTGTTGTCACTGCATTG-3'
β8 Y172M F	5'-GATTCATAATCAATGCAGTGACATGAATTTAGACTGCATGCC-3'
β8 Y172M R	5'-GGCATGCAGTCTAAATTCATGTCACTGCATTGATTATGAATC-3'
Y172A F	5'-GATTCATAATCAATGCAGTGACGCCAATTTAGACTGCATGCC-3'
Y172A R	5'-GGCATGCAGTCTAAATTGGCGTCACTGCATTGATTATGAATC-3'
β6 I183N F	5'-CCCTTGCAGTAGTAATCCATACTTCTG-3'
β6 I183N R	5'-CAGAAGTATGGATTACTACTGCAAGGG-3'
Porcine L-TGF-β1 C4S RGE F	5'-CCGCCGGGGTGAACTGGCCAC-3'
Porcine L-TGF-β1 C4S RGE R	5'-GTGGCCAGTTCACCCCGGCGG-3'
Porcine L-TGF-β1 C4S R249A F	5'-ACCTGCACAGCTCCCGGCACCGCGCAGCCCTGG-3'.
Porcine L-TGF-β1 C4S R249A R	5'-GTGGCCAGTTCACCCCGGCGG-3'
Human L-TGF-β1 F	5'-ATGGCCACCCCGCTGG-3'
Human L-TGF-β1 R	5'-CTCTACTAGTCTCGAGTTATCAGCTGCACTTGCAGGAGCGCAC-3'.
Puromycin cassette F	5'-ATCGTTTCAGACCCACCTCCC-3'
Puromycin cassette R	5'-CTCTGCTTAGCGAATTCGTTAACTGGCACCGGG-3'
Human L-TGF-β1 R249A F	5'-CACCGCGCAGCCCTGGACACCAAC-3'
Human L-TGF-β1 R249A R	5'-CCAGGGCTGCGCGGTGCCGGGAG-3'

Table S5 Primer sequences, related to Biologic Resources Table